

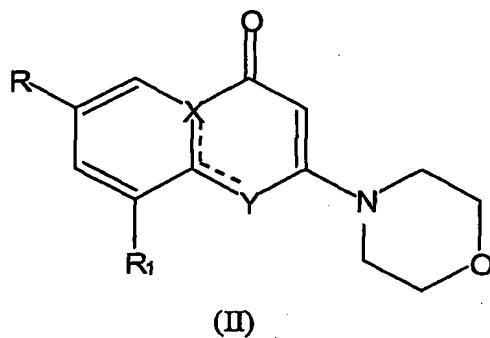
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WE CLAIM:

1. A method of disrupting platelet aggregation and adhesion occurring under high shear conditions comprising administering an effective amount of a selective PI 3-kinase inhibitor to a patient in need thereof.
2. A method for antithrombosis comprising administering an effective amount of a selective PI 3-kinase β inhibitor to a patient in need thereof, provided that the inhibitor is not according to formula (II):



wherein,

R is H, OH, F, Cl, Br, I, C_1 - C_6 alkyl, aryl or $(CH_2)_n$ -aryl;
 R^1 is H, OH, F, Cl, Br, I, C_1 - C_6 alkyl, C_3 - C_6 cycloalkyl, $CH=CH$ -aryl, $C\equiv C$ -aryl, $(CHR^3)_n$ -aryl, NR^3 - C_1 - C_6 alkyl, NR^3 -cycloalkyl, NR^3 - $(CHR^3)_n$ -aryl, $(CHR^3)_n$ - NR^3 -alkyl, $(CHR^3)_n$ - NR^3 -cycloalkyl, $(CHR^3)_n$ -O-aryl, $(CHR^3)_n$ -O-alkyl, $(CHR^3)_n$ -O-cycloalkyl, O- $(CHR^3)_n$ -aryl, S- $(CHR^3)_n$ -aryl, or CO-aryl, wherein n is 0, 1, or 2 and alkyl, cycloalkyl or aryl is optionally substituted with F, Cl, Br, I, CN, CO_2H , CO_2R^3 , NO_2 , CF_3 , substituted or unsubstituted C_1 - C_6 alkyl, substituted or unsubstituted cycloalkyl, substituted or unsubstituted aryl, OCF_3 , OR^3 , OSO_2 -aryl, substituted or unsubstituted amine, $NHCOR^3$, $NHSO_2R^3$, $CONHR^3$, or SO_2NHR^3 ;

and

R^3 is H, or substituted or unsubstituted C_1 - C_6 alkyl, substituted or unsubstituted aryl; except where the compound of formula (II) is selected from the group consisting of: 9-(3-pyridinylmethyl)oxy-2-morpholinyl-4H-pyrido[1,2-a]pyrimidin-4-one (TGX-140);

7-methyl-9-phenylaminomethyl-2-morpholinyl-4H-pyrido[1,2-a]pyrimidin-4-one (TGX-183);

8-(4-methylphenyl)2-morpholinyl-4(1H)-quinolinone (TGX-113);

8-(4-fluorophenoxy)-2-(4-morpholinyl)-4(1H)-quinolinone (TGX-121);

2-morpholinyl-8-(phenylmethyl)-4H-1-benzopyran-4-one (TGX-90);

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2-(4-morpholinyl)-8-(4-fluoro-2-methylphenyl)oxy-4H-1-benzopyran-4-one (TGX-184);

9-[(2-chlorophenyl)-methyl]amino-7-methyl-2-(4-morpholinyl)-4H-pyrido[1,2-a]pyrimidin-4-one (TGX-167);

9-[(2-methoxyphenyl)-methyl]amino]-7-methyl-2-(4-morpholinyl)-4H-pyrido[1,2-a]pyrimidin-4-one (TGX-137);

7-methyl-2-(4-morpholinyl)-9-[(phenylmethyl)amino]-4H-pyrido[1,2-a]pyrimidin-4-one (TGX-126);

9-[(4-fluoro-2-methylphenyl)amino]-7-methyl-2-(4-morpholinyl)-4H-pyrido[1,2-a]pyrimidin-4-one (TGX-170);

7-methyl-2-(4-morpholinyl)-9-[(1R)-1-phenylethyl]amino]-4H-pyrido[1,2-a]pyrimidin-4-one (TGX-123);

7-methyl-2-(4-morpholinyl)-9-[(2-pyridinylmethyl)amino]-4H-pyrido[1,2-a]pyrimidin-4-one (TGX-161);

9-[(4-chlorophenyl)methyl]amino]-7-methyl-2-(4-morpholinyl)-4H-pyrido[1,2-a]pyrimidin-4-one (TGX-108);

2-(4-morpholinyl)-9-(phenylmethyl)-4H-pyrido[1,2-a]pyrimidin-4-one (TGX-040);

7-methyl-9-(N-Methyl-N-phenyl)aminomethyl-2-(4-morpholinyl)-4H-pyrido[1,2-a]pyrimidin-4-one (TGX-195);

2-(4-morpholinyl)-8-(phenylmethyl)oxy-4H-1-benzopyran-4-one (TGX-102);

2-(4-morpholinyl)-8-(phenylmethyl)amino-4H-1-benzopyran-4-one (TGX-204);

2-(4-morpholinyl)-8-phenylamino-4H-1-benzopyran-4-one (TGX-324);

8-(3-chlorophenyl)oxy-2-(4-morpholinyl)-4H-1-benzopyran-4-one (TGX-259);

8-(3-methylphenyl)-2-(4-morpholinyl)-4(1H)-quinolinone (TGX-127);

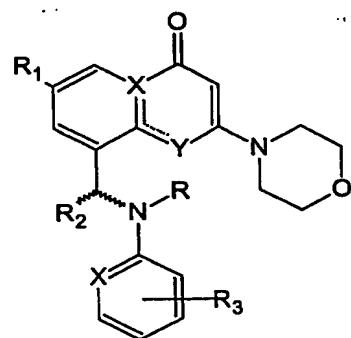
8-(2-fluorophenyl)-2-(4-morpholinyl)-4(1H)-quinolinone (TGX-143);

(\pm)-7-methyl-2-morpholin-4-yl-9-[1-(3-pyridinylamino)ethyl]-4H-pyrido[1,2-a]pyrimidin-4-one (KN-304).

3. The method of claim 2, wherein the selective PI 3-kinase β inhibitor is according to formula (I):

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(I)

wherein,

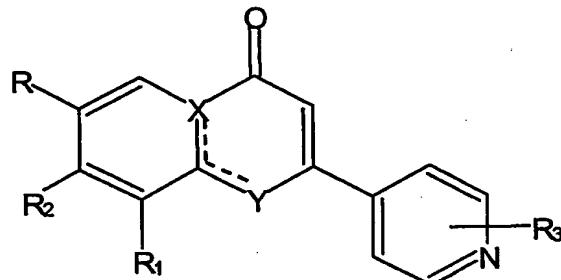
R is H, C₁-C₆ branched or straight chain alkyl, or aryl or (CH₂)_n-aryl;R₁ is H, OH, OCH₃, OCF₃, F, Cl, CF₃, C₁-C₆ branched or straight chain alkyl, or aryl or (CH₂)_n-aryl;R₂ is H, C₁-C₆ branched or straight chain alkyl, or aryl or (CH₂)_n-aryl in either the R

or the S configuration

R₃ is one or more of H, F, Cl, Br, I, CN, CO₂H, CO₂R, NO₂, CF₃, substituted or unsubstituted C₁-C₆ alkyl, substituted or unsubstituted cycloalkyl, substituted or unsubstituted aryl, OCH₃, OCH₂F, OCHF₂, OCF₃, OR, OSO₂-aryl, substituted or unsubstituted amine, NHCOR, NHSO₂R, CONHR, or SO₂NHR

X is C or N and Y is N or O.

4. The method of claim 2, wherein the selective PI 3-kinase β inhibitor is according to formula (III):



(III)

where X and Y are C and O respectively, or C and NH respectively, or both N.

R is H, OH, OCH₃, OCF₃, F, Cl, Br, I, C₁-C₆ alkyl, aryl or (CH₂)_n-aryl;R₁, R₂ and R₃ are independently H, OH, F, Cl, Br, I, C₁-C₆ alkyl, C₃-C₆ cycloalkyl, CH=CH-aryl, C≡C-aryl, (CHR')_n-aryl, NR'-(C₁-C₆ alkyl), NR'-(C₃-C₆ cycloalkyl), NR'-(CHR')_n-

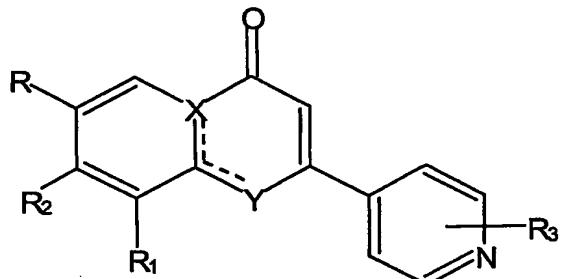
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aryl, $(CHR'^3)_n$ -NR 3* -aryl, $(CHR'^3)_n$ -NR 3* -alkyl, $(CHR'^3)_n$ -NR 3* -cycloalkyl, $(CHR'^3)_n$ -O-aryl, $(CHR'^3)_n$ -O-alkyl, $(CHR'^3)_n$ -O-cycloalkyl, O-($CHR'^3)_n$ -aryl, S-($CHR'^3)_n$ -aryl, or CO-aryl, wherein n is 0, 1, or 2 and alkyl, cycloalkyl or aryl is optionally substituted with F, Cl, Br, I, CN, CO₂H, CO₂R 3* , NO₂, CF₃, substituted or unsubstituted C₁-C₆ alkyl, substituted or unsubstituted cycloalkyl, substituted or unsubstituted aryl, OCF₃, OR 3* , OSO₂-aryl, substituted or unsubstituted amine, NHCOR 3* , NHSO₂R 3* , CONHR 3* , or SO₂NHR 3* ; and R 3* is H, or substituted or unsubstituted C₁-C₆ alkyl, substituted or unsubstituted aryl.

5. A compound having the following formula (III):



(III)

where X and Y are C and O respectively, or C and NH respectively, or both N.

R is H, OH, OCH₃, OCF₃, F, Cl, Br, I, C₁-C₆ alkyl, aryl or $(CH_2)_n$ -aryl;

R₁, R₂ and R₃ are independently H, OH, F, Cl, Br, I, C₁-C₆ alkyl, C₃-C₆ cycloalkyl, CH=CH-aryl, C≡C-aryl, $(CHR'^3)_n$ -aryl, NR 3* -C₁-C₆ alkyl, NR 3* -cycloalkyl, NR 3* - $(CHR'^3)_n$ -aryl, $(CHR'^3)_n$ -NR 3* -aryl, $(CHR'^3)_n$ -NR 3* -alkyl, $(CHR'^3)_n$ -NR 3* -cycloalkyl, $(CHR'^3)_n$ -O-aryl, $(CHR'^3)_n$ -O-alkyl, $(CHR'^3)_n$ -O-cycloalkyl, O-($CHR'^3)_n$ -aryl, S-($CHR'^3)_n$ -aryl, or CO-aryl, wherein n is 0, 1, or 2 and alkyl, cycloalkyl or aryl is optionally substituted with F, Cl, Br, I, CN, CO₂H, CO₂R 3* , NO₂, CF₃, substituted or unsubstituted C₁-C₆ alkyl, substituted or unsubstituted cycloalkyl, substituted or unsubstituted aryl, OCF₃, OR 3* , OSO₂-aryl, substituted or unsubstituted amine, NHCOR 3* , NHSO₂R 3* , CONHR 3* , or SO₂NHR 3* ; and

R 3* is H, or substituted or unsubstituted C₁-C₆ alkyl, substituted or unsubstituted aryl.

6. The method of claim 2, comprising administering the 2-morpholino-substituted derivative of formula (I) wherein:

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R is H, C₁-C₆ branched or straight chain alkyl or aryl;
R₁ is H, OH, OCH₃, OCF₃, F, Cl, CF₃, C₁-C₆ branched or straight chain alkyl;
R₂ is H, C₁-C₆ branched or straight chain alkyl, or aryl in either the R or the S configuration

R₃ is one or more of H, F, Cl, Br, CN, CO₂H, CO₂R, NO₂, CF₃, branched or straight chain C₁-C₆ alkyl, substituted or unsubstituted cycloalkyl, substituted or unsubstituted aryl, OCH₃, OCH₂F, OCHF₂, OCF₃, OR, substituted or unsubstituted amine, NHCOR, NHSO₂R, CONHR, or SO₂NHR

X is C or N and Y is N or O.

7. The method of claim 2, wherein the inhibitor administered is selected from the group consisting of:

(±)-7-methyl-9-{[methyl(phenyl)amino]methyl}-2-morpholin-4-yl-pyrido[1,2-a]pyrimidin-4-one (TGX-195);
(±)-7-methyl-2-morpholin-4-yl-9-(1-phenylaminoethyl)-pyrido[1,2-a]pyrimidin-4-one (TGX-221);
(±)-7-methyl-2-morpholin-4-yl-9-[1-(4-fluorophenylamino)ethyl]-pyrido[1,2-a]pyrimidin-4-one (TGX-224);
(±)-9-[1-(3,4-difluorophenylamino)ethyl]-7-methyl-2-morpholin-4-yl-pyrido[1,2-a]pyrimidin-4-one (TGX-237);
(±)-9-[1-(2,5-difluorophenylamino)ethyl]-7-methyl-2-morpholin-4-yl-pyrido[1,2-a]pyrimidin-4-one (TGX-238);
(±)-9-[1-(3,5-difluorophenylamino)ethyl]-7-methyl-2-morpholin-4-yl-pyrido[1,2-a]pyrimidin-4-one (TGX-239);
(±)-9-[1-(4-fluoro-2-methylphenylamino)ethyl]-7-methyl-2-morpholin-4-yl-pyrido[1,2-a]pyrimidin-4-one (TGX-240);
(±)-9-[1-(4-chlorophenylamino)ethyl]-7-methyl-2-morpholin-4-yl-pyrido[1,2-a]pyrimidin-4-one (TGX-243);
(±)-9-[1-(3,4-dichlorophenylamino)ethyl]-7-methyl-2-morpholin-4-yl-pyrido[1,2-a]pyrimidin-4-one (TGX-244);
(±)-9-[1-(3fluorophenylamino)ethyl]-7-methyl-2-morpholin-4-yl-pyrido[1,2-a]pyrimidin-4-one (TGX-247);
(±)-9-[1-(3-chlorophenylamino)ethyl]-7-methyl-2-morpholin-4-yl-pyrido[1,2-a]pyrimidin-4-one (TGX-248);

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(\pm)-7-methyl-2-morpholin-4-yl-9-[1-(2-thiazolylamino)ethyl]-pyrido[1,2-a]pyrimidin-4-one (TGX-261);

(\pm)-7-methyl-9-[1-(3-methylphenylamino)ethyl]-2-morpholin-4-yl-pyrido[1,2-a]pyrimidin-4-one (TGX-262);

(\pm)-7-methyl-2-morpholin-4-yl-9-[1-(3-trifluoromethylphenylamino)ethyl]-pyrido[1,2-a]pyrimidin-4-one (TGX-264); and

(\pm)-7-methyl-2-morpholin-4-yl-9-[1-(2-pyridinylamino)ethyl]-pyrido[1,2-a]pyrimidin-4-one (TGX-295).

(\pm)-2-({1-[7-methyl-2-(morpholin-4-yl)-4-oxo-pyrido[1,2-a]pyrimidin-9-yl]ethyl}amino)benzoic acid (KN-309);

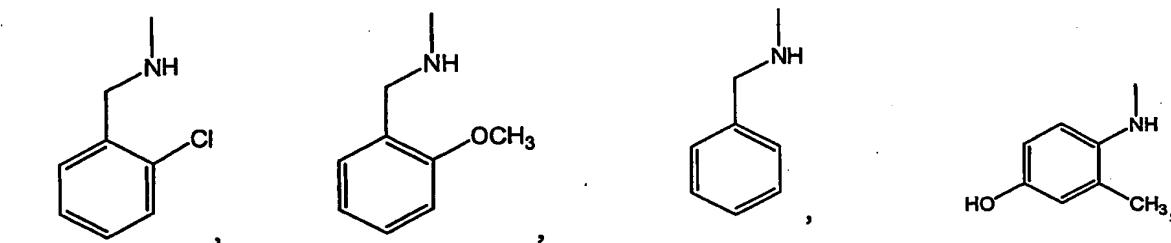
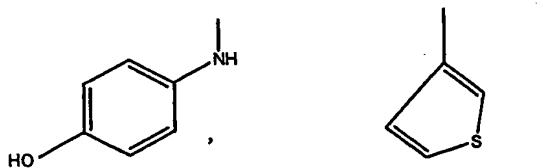
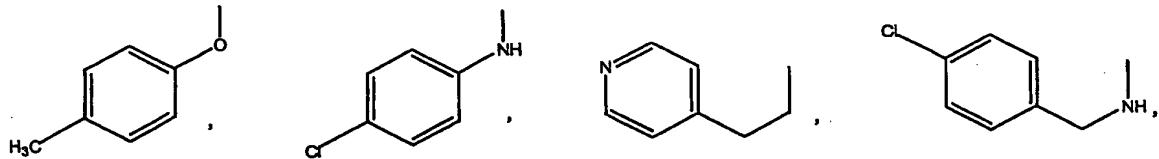
(\pm) methyl 2-({1-[7-methyl-2-(morpholin-4-yl)-4-oxo-pyrido[1,2-a]pyrimidin-9-yl]ethyl}amino)benzoate (KN-321);

(\pm)-2-({1-[7-methyl-2-(morpholin-4-yl)-4-oxo-pyrido[1,2-a]pyrimidin-9-yl]ethyl}amino)benzonitrile (KN-320);

(\pm)-7-methyl-2-(morpholin-4-yl)-9-(1-{{2-(2H-tetrazol-5-yl)phenyl}amino}ethyl)-pyrido[1,2-a]pyrimid-4-one (KN-325);

(\pm)-2-(4-morpholinyl)-8[1-(phenylamino)ethyl]-4H-1-benzopyran-4-one (TGX-280).

8. The compound of claim 5, wherein R¹ is selected from a group consisting of, CH₃, C₂H₅,

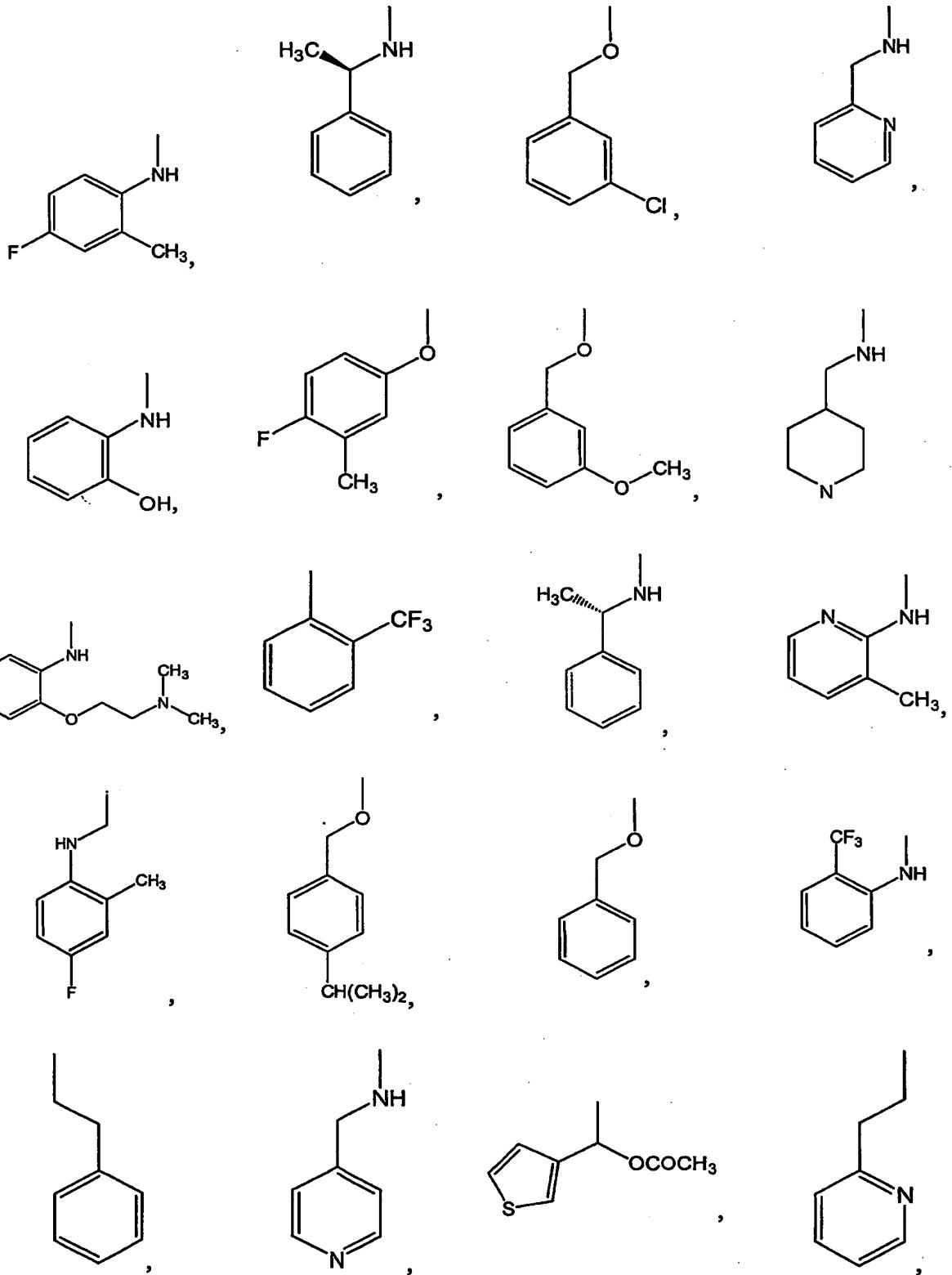


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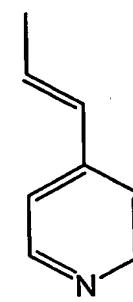
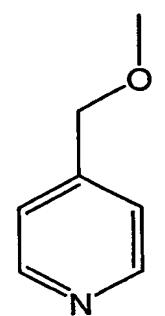
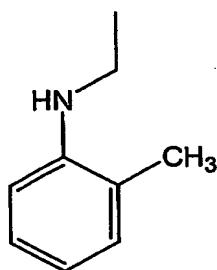
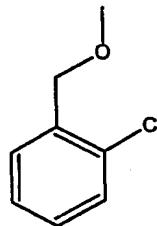
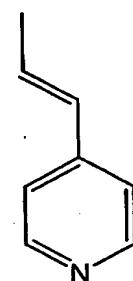
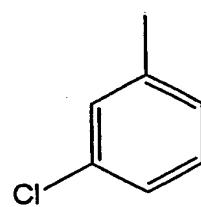
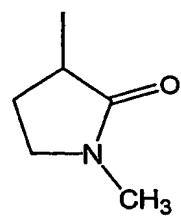
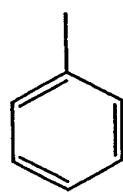
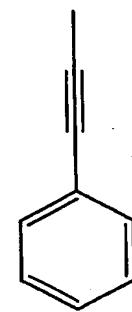
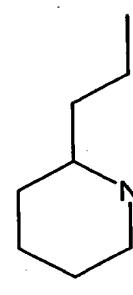
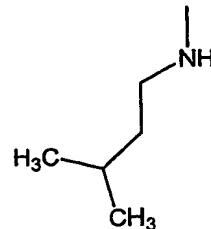
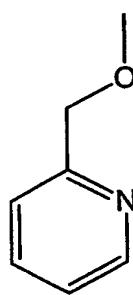
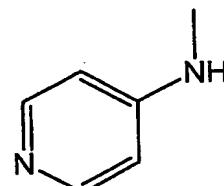
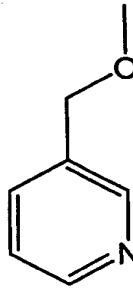
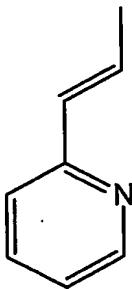
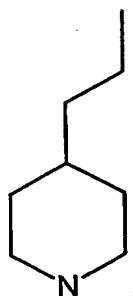


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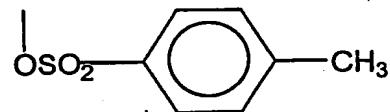
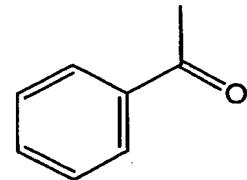
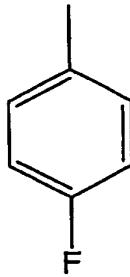
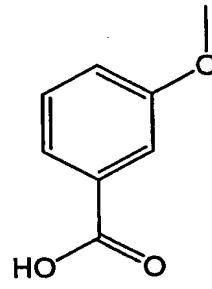
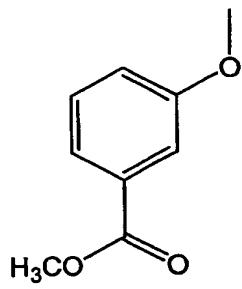
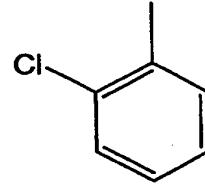
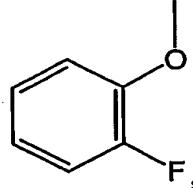
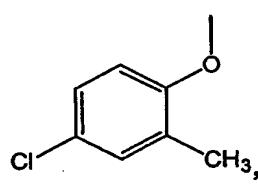
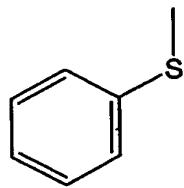
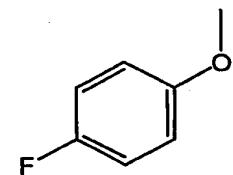
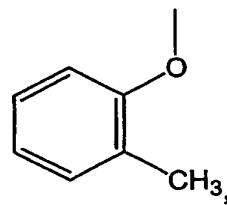
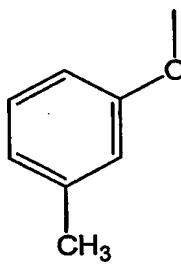
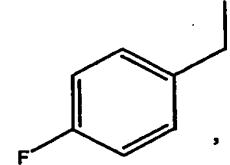
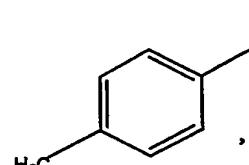
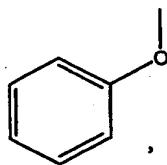
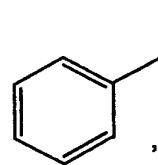


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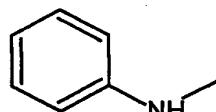
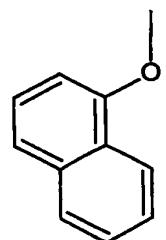
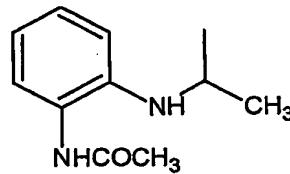
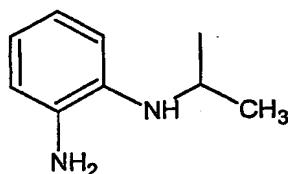
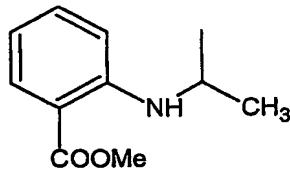
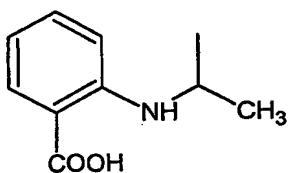
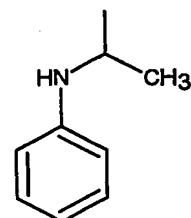
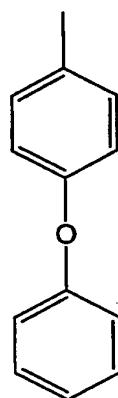
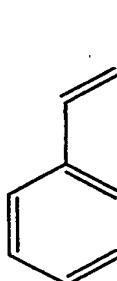
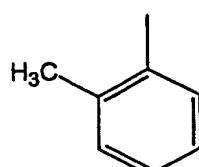
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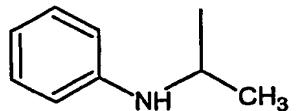
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9. The compound of claim 5, wherein R is methyl and R¹ is

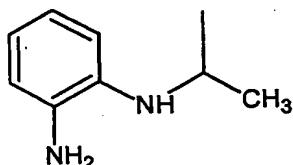


10. The compound of claim 5, wherein R is methyl and R¹ is

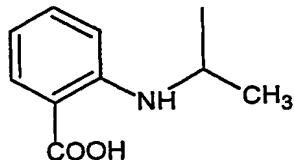
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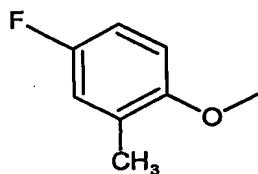
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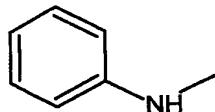
11. The compound of claim 5, wherein R is methyl and R¹ is



12. The compound of claim 5, wherein R is H and R¹ is



13. The compound of claim 5, wherein R is H and R¹ is



14. A method for inhibiting phosphoinositide 3-kinase in a patient, comprising administering to a patient an amount of the compound of claim 5 effective in inhibiting the phosphoinositide 3-kinase in the patient.

15. A method for preventing or treating cardiovascular disease comprising administering an effective amount of the compound of claim 5 to a patient in need thereof.

16. A method for preventing or treating respiratory disease comprising administering an effective amount of the compound of claim 5 to a patient in need thereof.

17. A method for preventing or treating cancer comprising administering an effective amount of the compound of claim 5 to a patient in need thereof.

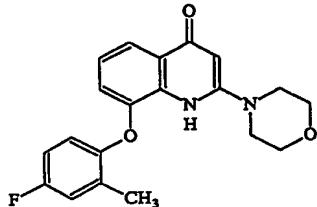
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18. A method for preventing or treating disease linked to disordered white blood cell function comprising administering an effective amount of the compound of claim 5 to a patient in need thereof.

19. The method of claim 2, wherein the inhibitor administered is:



20. The method of claim 4, wherein the inhibitor administered is 6-methyl-8-[1-(phenylamino)ethyl]-2-(4-pyridinyl)-4H-benzopyran-4-one.

21. The method of claim 4, wherein the inhibitor administered is 6-methyl-8-{1-[(2-aminophenyl)amino]ethyl}-2-(4-pyridinyl)-4H-benzopyran-4-one.

22. A compound which is (\pm)-7-methyl-2-morpholin-4-yl-9-(1-phenylaminoethyl)-pyrido[1,2-a]pyrimidin-4-one.

23. A compound which is (\pm)-2-({1-[7-methyl-2-(morpholin-4-yl)-4-oxo-pyrido[1,2-a]pyrimidin-9-yl]ethyl} amino)benzoic acid.

24. A compound which is (\pm)-2-({1-[7-methyl-2-(morpholin-4-yl)-4-oxo-pyrido[1,2-a]pyrimidin-9-yl]ethyl} amino)benzonitrile.

25. A compound which is (\pm) methyl 2-({1-[7-methyl-2-(morpholin-4-yl)-4-oxo-pyrido[1,2-a]pyrimidin-9-yl]ethyl} amino)benzoate.

26. A compound which is (\pm)-7-methyl-2-(morpholin-4-yl)-9-(1-{{2-(2H-tetrazol-5-yl)phenyl}amino}ethyl)-pyrido[1,2-a]pyrimid-4-one.